PATENT COOPERATION TREATY

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From the INTERNATIONAL SEARCHING AUTHORITY

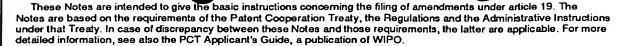
NOTIFICATION OF TRANSMITTAL OF GLAXO SMITH KLINE THE INTERNATIONAL SEARCH REPORT Attn. Levy, David J. OR THE DECLARATION Five Moore Drive PO Box 13398 (PCT Rule 44.1) Research Triangle Park, NC 27709 UNITED STATES OF AMERICA Date of mailing (day/month/year) 15/04/2004 Applicant's or agent's file reference FOR FURTHER ACTION See paragraphs 1 and 4 below PU4963WO International application No. International filing date (day/month/year) 12/12/2003 PCT/US 03/39975 Applicant SMITHKLINE BEECHAM CORPORATION

1.	\mathbf{x}	The app	licant is hereby r	notified that the International Search Report has been established and is transmitted herewith.			
				and statement under Article 19: if he so wishes, to amend the claims of the International Application (see Rule 46):			
		When? The time limit for filing such amendments is normally 2 months from the date of transmittal of the International Search Report; however, for more details, see the notes on the accompanying sheet.					
	٠	Where?	Directly to the	International Bureau of WIPO 34, chemin des Colombettes 1211 Geneva 20, Switzerland Fascimile No.: (41–22) 740.14.35			
		For mor	e detailed instr	uctions, see the notes on the accompanying sheet.			
2.				notified that no International Search Report will be established and that the declaration under ect is transmitted herewith.			
3.		With reg	ard to the prote	est against payment of (an) additional fee(s) under Rule 40.2, the applicant is notified that:			
				with the decision thereon has been transmitted to the International Bureau together with the to forward the texts of both the protest and the decision thereon to the designated Offices.			
		no	decision has bee	en made yet on the protest; the applicant will be notified as soon as a decision is made.			
4.	Furti	ner actio	n(s): The appl	icant is reminded of the following:			
	If the	né applica prity claim	ant wishes to avo	he priority date, the international application will be published by the International Bureau. oid or postpone publication, a notice of withdrawal of the international application, or of the International Bureau as provided in Rules 90 <i>bis</i> .1 and 90 <i>bis</i> .3, respectively, before the reparations for international publication.			
,				ority date, a demand for international preliminary examination must be filed if the applicant into the national phase until 30 months from the priority date (in some Offices even later).			
,	bef	ore all de	signated Offices	ority date, the applicant must perform the prescribed acts for entry into the national phase which have not been elected in the demand or in a later election within 19 months from the elected because they are not bound by Chapter II.			

Name and mailing address of the International Searching Authority Authorized officer European Patent Office, P.B. 5818 Patentlaan 2 NL-2280 HV Rijswijk Tel. (+31-70) 340-2040, Tx. 31 651 epo nl, Fax: (+31-70) 340-3016 Angela Lopez Nava APR 2 2 2004 GLOSALINTTH ECTOM FROM PRINTRY

Form PCT/ISA/220 (July 1998)

NOTES TO FORM PCT/ISA/220



In these Notes, "Article", "Rule", and "Section" refer to the provisions of the PCT, the PCT Regulations and the PCT Administrative Instructions respectively.

INSTRUCTIONS CONCERNING AMENDMENTS UNDER ARTICLE 19

The applicant has, after having received the international search report, one opportunity to amend the claims of the international application. It should however be emphasized that, since all parts of the international application (claims, description and drawings) may be amended during the international preliminary examination procedure, there is usually no need to file amendments of the claims under Article 19 except where, e.g. the applicant wants the latter to be published for the purposes of provisional protection or has another reason for amending the claims before international policiation. Furthermore, it should be emphasized that provisional protection is available in some States only.

What parts of the international application may be amended?

Under Article 19, only the claims may be amended.

During the international phase, the claims may also be amended (or further amended) under Article 34 before the International Preliminary Examining Authority. The description and drawings may only be amended under Article 34 before the International Examining Authority.

Upon entry into the national phase, all parts of the international application may be amended under Article 28 or, where applicable, Article 41.

When?

Within 2 months from the date of transmittal of the international search report or 16 months from the priority date, whichever time limit expires later. It should be noted, however, that the amendments will be considered as having been received on time if they are received by the International Bureau after the expiration of the applicable time limit but before the completion of the technical preparations for international publication (Rule 46.1).

Where not to file the amendments?

The amendments may only be filed with the International Bureau and not with the receiving Office or the International Searching Authority (Rule 46.2).

Where a demand for international preliminary examination has been is filed, see below.

How?

Either by cancelling one or more entire claims, by adding one or more new claims or by amending the text of one or more of the claims as filed.

A replacement sheet must be submitted for each sheet of the claims which, on account of an amendment or amendments, differs from the sheet originally filed.

All the claims appearing on a replacement sheet must be numbered in Arabic numerals. Where a claim is cancelled, no renumbering of the other claims is required. In all cases where claims are renumbered, they must be renumbered consecutively (Administrative Instructions, Section 205(b)).

The amendments must be made in the language in which the international application is to be published.

What documents must/may accompany the amendments?

Letter (Section 205(b)):

The amendments must be submitted with a letter.

The letter will not be published with the international application and the amended claims. It should not be confused with the "Statement under Article 19(1)" (see below, under "Statement under Article 19(1)").

The letter must be in English or French, at the choice of the applicant. However, if the language of the international application is English, the letter must be in English; if the language of the international application is French, the letter must be in French.

NOTES TO FORM PCT/ISA/220 (continued)





The letter must indicate the differences between the claims as filed and the claims as amended. It must, in particular, indicate, in connection with each claim appearing in the international application (it being understood that identical indications concerning several claims may be grouped), whether

- (i) the claim is unchanged;
- (ii) the claim is cancelled;
- (iii) the claim is new:
- (iv) the claim replaces one or more claims as filed;
- (v) the claim is the result of the division of a claim as filed.

The following examples illustrate the manner in which amendments must be explained in the accompanying letter:

- [Where originally there were 48 claims and after amendment of some claims there are 51]:
 "Claims 1 to 29, 31, 32, 34, 35, 37 to 48 replaced by amended claims bearing the same numbers;
 claims 30, 33 and 36 unchanged; new claims 49 to 51 added."
- [Where originally there were 15 claims and after amendment of all claims there are 11]: "Claims 1 to 15 replaced by amended claims 1 to 11."
- 3. [Where originally there were 14 claims and the amendments consist in cancelling some claims and in adding new claims]:
 "Claims 1 to 6 and 14 unchanged; claims 7 to 13 cancelled; new claims 15, 16 and 17 added." or
 "Claims 7 to 13 cancelled; new claims 15, 16 and 17 added; all other claims unchanged."
- 4. [Where various kinds of amendments are made]: "Claims 1-10 unchanged; claims 11 to 13, 18 and 19 cancelled; claims 14, 15 and 16 replaced by amended claim 14; claim 17 subdivided into amended claims 15, 16 and 17; new claims 20 and 21 added."

"Statement under article 19(1)" (Rule 46.4)

The amendments may be accompanied by a statement explaining the amendments and indicating any impact that such amendments might have on the description and the drawings (which cannot be amended under Article 19(1)).

The statement will be published with the international application and the amended claims.

It must be in the language in which the international appplication is to be published.

It must be brief, not exceeding 500 words if in English or if translated into English.

It should not be confused with and does not replace the letter indicating the differences between the claims as filed and as amended. It must be filed on a separate sheet and must be identified as such by a heading, preferably by using the words "Statement under Article 19(1)."

It may not contain any disparaging comments on the international search report or the relevance of citations contained in that report. Reference to citations, relevant to a given claim, contained in the international search report may be made only in connection with an amendment of that claim.

Consequence if a demand for international preliminary examination has already been filed

If, at the time of filing any amendments under Article 19, a demand for international preliminary examination has already been submitted, the applicant must preferably, at the same time of filing the amendments with the International Bureau, also file a copy of such amendments with the International Preliminary Examining Authority (see Rule 62.2(a), first sentence).

Consequence with regard to translation of the international application for entry into the national phase

The applicant's attention is drawn to the fact that, where upon entry into the national phase, a translation of the claims as amended under Article 19 may have to be furnished to the designated/elected Offices, instead of, or in addition to, the translation of the claims as filed.

For further details on the requirements of each designated/elected Office, see Volume II of the PCT Applicant's Guide

PATENT COOPERATION TREATY



PCT



INTERNATIONAL SEARCH REPORT

(PCT Article 18 and Rules 43 and 44)

Applicant's or agent's file reference		ion of Transmittal of International Search Report SA/220) as well as, where applicable, item 5 below.
PU4963WO	ACTION	SW220) as well as, where applicable, item 3 below.
International application No.	International filing date (day/month/year	(Earliest) Priority Date (day/month/year)
PCT/US 03/39975	12/12/2003	13/12/2002
Applicant		
·		
SMITHKLINE BEECHAM CORPORA	ATION	
This International Search Report has been according to Article 18. A copy is being tra		Authority and is transmitted to the applicant
This International Search Report consists	of a total of5 sheets.	
X It is also accompanied by	a copy of each prior art document cited in	this report.
Basis of the report		·-
a. With regard to the language, the	international search was carried out on the ess otherwise indicated under this item.	e basis of the international application in the
the international search w. Authority (Rule 23.1(b)).	as carried out on the basis of a translation	of the international application furnished to this
b. With regard to any nucleotide and was carried out on the basis of the		ne international application, the international search
	nal application in written form.	
filed together with the inte	rnational application in computer readable	form.
furnished subsequently to	this Authority in written form.	
furnished subsequently to	this Authority in computer readble form.	
the statement that the sub international application as		ng does not go beyond the disclosure in the
the statement that the info furnished	rmation recorded in computer readable fo	rm is identical to the written sequence listing has been
2. X Certain claims were four	nd unsearchable (See Box I).	
3. Unity of invention is lack	king (see Box II).	
4. With regard to the title,		
X the text is approved as sul	omitted by the applicant.	
the text has been establish	ned by this Authority to read as follows:	
·		
5. With regard to the abstract,		
X the text is approved as sub	omitted by the applicant.	
the text has been establish	ned, according to Rule 38.2(b), by this Aut date of mailing of this international search	hority as it appears in Box III. The applicant may, a report, submit comments to this Authority.
6. The figure of the drawings to be public	shed with the abstract is Figure No.	
as suggested by the applic	eant.	None of the figures.
because the applicant faile	ed to suggest a figure.	
because this figure better of	characterizes the invention.	

INTERNATIONAL SEARCH REPORT

International Application No PCT/US 03/39975

A. CLASSIFICATION OF SUBJECT MATERIAL TO THE PROPERTY OF SUBJECT M CO7D413/0 211/58 CO7D471/10 A61K31/439 A61P31/18 According to International Patent Classification (IPC) or to both national classification and IPC **B. FIELDS SEARCHED** Minimum documentation searched (classification system followed by classification symbols) IPC 7 CO7D A61K Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched Electronic data base consulted during the international search (name of data base and, where practical, search terms used) EPO-Internal, WPI Data, BEILSTEIN Data, CHEM ABS Data C. DOCUMENTS CONSIDERED TO BE RELEVANT Citation of document, with indication, where appropriate, of the relevant passages Relevant to claim No. Category ' Α US 6 096 780 A (SHIRAISHI MITSURU ET AL) 1 - 391 August 2000 (2000-08-01) claim 1 Ε WO 2004/009584 A (AHN YOUNG GIL; HAM 1 YOUNG JIN (KR); KIM HAN KYONG (KR); KIM HEE SEOCK) 29 January 2004 (2004-01-29) claim 1; examples 1-8 Patent family members are listed in annex. Further documents are listed in the continuation of box C. Special categories of cited documents: "T" later document published after the international filing date or priority date and not in conflict with the application but cited to understand the principle or theory underlying the *A* document defining the general state of the art which is not considered to be of particular relevance invention earlier document but published on or after the international "X" document of particular relevance; the claimed invention cannot be considered novel or cannot be considered to involve an inventive step when the document is taken alone *L* document which may throw doubts on priority claim(s) or which is cited to establish the publication date of another "Y" document of particular relevance; the claimed invention citation or other special reason (as specified) cannot be considered to involve an inventive step when the document is combined with one or more other such docu-"O" document referring to an oral disclosure, use, exhibition or ments, such combination being obvious to a person skilled other means document published prior to the international filing date but later than the priority date claimed "&" document member of the same patent family Date of the actual completion of the international search Date of mailing of the international search report 30 March 2004 15/04/2004 Name and mailing address of the ISA Authorized officer European Patent Office, P.B. 5818 Patentlaan 2 NL - 2280 HV Rijswijk Tel. (+31-70) 340-2040, Tx. 31 651 epo nl, Seelmann, I Fax: (+31-70) 340-3016



Continuation of Box I.2

The present claims relate to an extremely large number of possible compounds. Support within the meaning of Article 6 PCT and/or disclosure within the meaning of Article 5 PCT is to be found, however, for only a very small proportion of the compounds claimed. In the present case, the claims so lack support, and the application so lacks disclosure, that a meaningful search over the whole of the claimed scope is impossible. Consequently, the search has been carried out for those parts of the claims which appear to be supported and disclosed, namely those parts relating to carbocyclic, bicyclic compounds. The following limitations were made in the search: All Z = carbon, B is a 4-7 membered saturated carbocyclic ring and R10 = R7 (two R10 cannot form a ring). All compounds of claim 22 are encompassed by the scope of the search.

The applicant's attention is drawn to the fact that claims relating to inventions in respect of which no international search report has been established need not be the subject of an international preliminary examination (Rule 66.1(e) PCT). The applicant is advised that the EPO policy when acting as an International Preliminary Examining Authority is normally not to carry out a preliminary examination on matter which has not been searched. This is the case irrespective of whether or not the claims are amended following receipt of the search report or during any Chapter II procedure. If the application proceeds into the regional phase before the EPO, the applicant is reminded that a search may be carried out during examination before the EPO (see EPO Guideline C-VI, 8.5), should the problems which led to the Article 17(2) declaration be overcome.

INTERNATIONAL SEARCH REPORT

International application No. PCT/US 03/39975

Box I	Observations where common claims were found unsearchable (Continua and of item 1 of first sheet)
This Inte	ernational Search Report has not been established in respect of certain claims under Article 17(2)(a) for the following reasons:
1.	Claims Nos.: because they relate to subject matter not required to be searched by this Authority, namely:
2. X	Claims Nos.: because they relate to parts of the International Application that do not comply with the prescribed requirements to such an extent that no meaningful International Search can be carried out, specifically: See FURTHER INFORMATION sheet PCT/ISA/210
3.	Claims Nos.: because they are dependent claims and are not drafted in accordance with the second and third sentences of Rule 6.4(a).
Box II	Observations where unity of invention is lacking (Continuation of item 2 of first sheet)
This Inte	rnational Searching Authority found multiple inventions in this international application, as follows:
1.	As all required additional search fees were timely paid by the applicant, this International Search Report covers all searchable claims.
2.	As all searchable claims could be searched without effort justifying an additional fee, this Authority did not invite payment of any additional fee.
	As only some of the required additional search fees were timely paid by the applicant, this International Search Report covers only those claims for which fees were paid, specifically claims Nos.:
4.	No required additional search fees were timely paid by the applicant. Consequently, this International Search Report is restricted to the invention first mentioned in the claims; it is covered by claims Nos.:
Remark (The additional search fees were accompanied by the applicant's protest. No protest accompanied the payment of additional search fees.

INTERNATIONAL SEARCH REPORT

Information on patent family members

International Application No PCT/US 03/39975

Patent document cited in search report		Publication date		Patent family member(s)		Publication date
US 6096780	A	01-08-2000	AU	748064	B2	30-05-2002
			ΑU	5301599	Α	14-03-2000
			BR	9813691	Α	10-10-2000
			CA	2304959	A1	01-07-1999
			CA	2337307	A1	02-03-2000
			CN	1282243	T	31-01-2001
			ΕP	1039899	A2	04-10-2000
			EP	1104416	A2	06-06-2001
			HU	0100199	A2	28-04-2001
			ID	29994	Α	25-10-2001
			WO	9932100	A2	01-07-1999
			WO	0010965	A2	02-03-2000
			JP	2000128842	Α	09-05-2000
			JP	2000128782	Α	09-05-2000
			US	6172061	B1	09-01-2001
			US	6376536	B1	23-04-2002
			US	6268354	B1	31-07-2001
WO 2004009584	Α	29-01-2004	WO	2004009584	A1	29-01-2004